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(A) 004/007

Docket No 930008-2200

## IN THE UNITED STATES PATENT AND TRADEMARK ()FFICE

In re Application of Gerhard Floefle et al.

Application No.: 10/535,630

Confirmation No.: 9251

Filed: October 20, 2003 (PCT/EP03/011603)

Art Unit: 1625

For: TUBULYSINS, PREPARATION PROCESSES

AND TUBULYSIN PREPARATIONS

Examiner: John Mabry

Commissioner for Patents P.O. Box 1450 Alexandrin, VA 22313-1450

## DECLARATION OF WOLFGANG RICHTER, PH.D. UNDER 37 C.F.R. 5 1,132

- I, Dr. Wolfgang Richter, declare as follows:
- 1. I am managing Director of R&D Biopharmacouticals GmbH, Munich, Germany. My field of research and expectise is in the area of organic and medicinal chemistry.
- 2. The following experimental data was obtained in accordance with the following citations: T.-T. Yang, P. Sinai, S.R. Kain, Analytical Biochemistry 241, 103-108 (1996); G. Kaur, M. Hollingshead, S. Holbeck, V. Schauer-Vukasinovic, R.F. Camalier, A. Dömling, S. Agarwal, Biochem. J. (2006) 396, 235-242.
- 3. The following experimental data shows that derivatives 2, 3, 5 and 6 carrying an ester group at residue R show a higher activity than derivatives 1 (tubulysin A) and 4 wherein R is OH.

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Application No. 10/535,630 Declaration of Wolfgang Richter, under 37 C.F.R. § 1.132

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AN: 0232365929

No	Structure	TURULYSIN	1.929 IC50 [ng/ml]
1		Tubulysin A	0.2
2		NT5	0.02
3		NT6	0,02
4		NT9	0.1
5		NT10	0.04
6		NT14	0.05

AN: 0232365929

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VON: R&D BIOPHARMACEUTICA 08970076499

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Application No. 10/535,630 Declaration of Wolfgang Richtor, under 37 C.F.R. § 1.132

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Docket No. 930008-2200

4. I heroby declare that all statements made herein of my own knowledge are two and that all statements made on information and belief are believed to be true; and, that these statements were made with the knowledge that willful false statements and the like so triade are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful statements may jeopardize the validity of the application or any patent issued thereon.

Date: May 18 14, 2003

Respectfully Submitted,

Dr. Wolf jany Richter Hunging Director R&D-Biophanna conticols Guns U